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81. A modified wild-type human thyroid stimulating hormone (TSH) having increased TSH activity compared to wild-type human TSH comprising an  $\alpha$ -subunit and a  $\beta$ -subunit, said  $\alpha$ -subunit comprising at least three basic amino acids in the  $\alpha$ -subunit at positions selected from the group consisting of positions 11, 13, 14,16, 17, and 20.

- 82. The modified wild-type human TSH of Claim 81, said  $\alpha$ -subunit further comprising a fourth basic amino acid at a position selected from the group consisting of positions 11, 13, 14, 16, 17, and 20.
- 83. The modified wild-type human TSH of Claim 82, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 11, 13, 16, and 20.
- 84. The modified wild-type human TSH of Claim 82, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 11, 13, 17, and 20.
- 85. The modified wild-type human TSH of Claim 82, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 13, 14, 16, and 20.
- 86. The modified wild-type human TSH of Claim 82, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 13, 14, 17, and 20.
- 87. The modified wild-type human TSH of Claim 82, said  $\alpha$ -subunit further comprising a fifth basic amino acid at a position selected from the group consisting of positions 11, 13, 14, 16, 17, and 20.
- 88. The modified wild-type human TSH of Claim 87, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 18, 14, 16, 17, and 20.
- 89. The modified wild-type human TSH of Claim 87, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 11, 13, 14, 16, and 20.
- 90. The modified wild-type human  $\frac{1}{3}$ SH of Claim 81, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 11, 13, 14, 16, 17, and 20.
- 91. The modified wild-type human TSH of Claim 81, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 13, 16, and 20.
- 92. The modified wild-type human TSH of Claim 81, further modified so that said  $\beta$ -subunit comprises a basic amino acid in the  $\beta$ -subunit in at least one position selected from the group consisting of positions 58, 63, and 69

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93. The modified wild-type human TSH of Claim 92, wherein the basic amino acids of the  $\beta$ -subunit are at positions 58, 63, and 69.

- 94. The modified wild-type human TSH of Claim 92, wherein a basic amino acid of the  $\beta$ -subunit is at position 58.
- 95. The modified wild-type human TSH of Claim 92, wherein a basic amino acid of the β-subunit is at position 63.
- 96. The modified wild-type human TSH of Claim 92, wherein a basic amino acid of the β-subunit is at position 69.
- 97. The modified wild type human TSH of Claim 81, wherein the basic amino acids are selected from the group consisting of lysine and arginine.
- 98. A nucleic acid encoding the modified wild-type human TSH  $\alpha$ -subunit of Claim 81.

1996. A vector comprising the nucleic acid of Claim 98, wherein the vector is suitable for expressing the nucleic acid.

20 100. A host cell comprising the vector of Claim 93, wherein the host cell is suitable for expressing the nucleic acid.

7 101. The modified wild-type human TSH of Claim 81, further modified so that said modified wild-type human TSH has less than five amino acid substitutions in said  $\alpha$ -subunit in positions other than positions 11, 13, 14, 16, 17, and 20.

- 102. The modified wild-type human TSH of Claim 81, further modified so that said modified wild-type human TSH has less than four amino acid substitutions in said  $\alpha$ -subunit in positions other than positions 11, 13, 14, 16, 17, and 20.
- 103. The modified wild-type human TSH of Claim 81, further modified so that said modified wild-type human TSH has less than three amino acid substitutions in said  $\alpha$ -subunit in positions other than positions 11, 13, 14, 16, 17, and 20.
- 104. The modified wild-type human TSH of Claim 81, further modified so that said modified wild-type human TSH has less than two amino acid substitutions in said  $\alpha$ -subunit in positions other than positions 11, 13, 14, 16, 17, and 20.
- 105. The modified wild-type human TSH of Claim 81, further modified so that said modified wild-type human TSH has complete amino acid sequence homology with

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the corresponding wild-type human TSH in said  $\alpha$ -subunit in positions other than positions 11, 13, 14, 16, 17, and 20.

- 106. A modified wild-type human thyroid stimulating hormone (TSH) having increased TSH activity compared to wild-type human TSH comprising an  $\alpha$ -subunit and a  $\beta$ -subunit, said  $\alpha$ -subunit comprising a basic amino acid in the  $\alpha$ -subunit in at least one position selected from the group consisting of positions 11, 13, 14,16, 17, and 20.
- 107. The modified wild-type human TSH of Claim 106, wherein a basic amino acid of the  $\alpha$ -subunit is at position 11.
- 108. The modified wild-type human TSH of Claim 106, wherein a basic amino acid of the  $\alpha$ -subunit is at position 13.
- 109. The modified wild-type human TSH of Claim 106, wherein a basic amino acid of the  $\alpha$ -subunit is at position 14.
- 110. The modified wild-type human TSH of Claim 106, wherein a basic amino acid of the  $\alpha$ -subunit is at position 16.
- 111. The modified wild-type human TSH of Claim 106, wherein a basic amino acid of the  $\alpha$ -subunit is at position  $\sqrt{7}$ .
- 112. The modified wild-type human TSH of Claim 106, wherein a basic amino acid of the  $\alpha$ -subunit is at position 20
- 113. The modified wild-type human TSH of Claim 106, wherein the basic amino acid is selected from the group consisting of lysine and arginine
- 114. The modified wild-type human TSH of Claim 106, further modified so that said  $\alpha$ -subunit comprises a basic amino acid in at least two positions selected from the group consisting of positions 11, 13, 14,16, 17, and 20.
- 115. The modified wild-type human  $\uparrow$ SH of Claim 114, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 16 and 20.
- 116. The modified wild-type human TSH of Claim 114, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 16 and 13.
- 117. The modified wild-type human TSH of Claim 114, wherein the basic amino acids of the  $\alpha$ -subunit are at positions 20 and 13.

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118. The modified wild-type human TSH of Claim 114, wherein the basic amino acid is selected from the group consisting of lysine and arginine.

- 119. The modified wild-type human TSH of Claim 106, further modified so that said  $\beta$ -subunit further comprises a basic amino acid in the  $\beta$ -subunit in at least one position selected from the group consisting of positions 58, 63, and 69.
- 120. The modified wild-type human TSH of Claim 119, wherein the basic amino acids of the β-subunit are at positions 58, 63, and 69.
- 121. The modified wild-type human TSH of Claim 119, wherein a basic amino acid of the β-subunit is at\position 58.
- 122. The modified wild-type human TSH of Claim 119, wherein a basic amino acid of the  $\beta$ -subunit is at position 63.
- 123. The modified wild type human TSH of Claim 119, wherein a basic amino acid of the β-subunit is at position 69.
- 124. A nucleic acid encoding the modified wild-type human TSH  $\alpha$ -subunit of Claim 106.

125. A vector comprising the nucleic acid of Claim124, wherein the vector is suitable for expressing the nucleic acid.

46126. A host cell comprising the vector of Claim 125, wherein the host cell is suitable for expressing the nucleic acid.

127. The modified wild-type human TSH of Claim 106, further modified so that said modified wild-type human TSH has less than five amino acid substitutions in said  $\alpha$ -subunit in positions other than positions 11, 13, 14, 16, 17, and 20.

128. The modified wild-type human TSH of Claim 106, further modified so that said modified wild-type human TSH has less than four amino acid substitutions in said  $\alpha$ -subunit in positions other than positions 11, 13, 14, 16, 17, and 20.

- 129. The modified wild-type human TSH of Claim 106, further modified so that said modified wild-type human TSH has less than three amino acid substitutions in said  $\alpha$ -subunit in positions other than positions 11, 13, 14, 16, 17, and 20.
- 130. The modified wild-type human TSH of Claim 106, further modified so that said modified wild-type human TSH has less than two amino acid substitutions in said  $\alpha$ -subunit in positions other than positions (1, 13, 14, 16, 17, and 20.

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